International application No. PCT/CA2005/000069

A. CLASSIFICATION OF SUBJECT MATTER

IPC7:A61K 48/00; A61P 35/00; A61K 31/7088; A61K 31/522; A61K 31/517; A61K 31/513

According to International Patent Classification (IPC) or to both national classification and IPC

### B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched IPC7:A61K 48/00; A61P 35/00; A61K 31/7088; A61K 31/522; A61K 31/517; A61K 31/513

Electronic database(s) consulted during the international search (name of database(s) and, where practicable, search terms used) Canadian Patent Database, Delphion, NCBI, PubMed, STN, QPAT Search terms: antisense, mesothelioma, thymidylate synthase, cancer

### C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 9963114 A1 (ISIS PHARMACEUTICALS, INC.) 09.12.1999 Whole document	18-23
Y	Whole document	1-17, 24-26
х	CA 2301957 A1 (ZENECA LIMITED, ISIS PHARMACEUTICALS) 01.04.1999 Whole document	18-23
Y		1-17, 24-26
Х	FERGUSON PJ et al Antisense down-regulation of thymidylate synthase to suppress growth and enhance cytotoxicity of 5-FUdR, 5-FU and Tomudex in HeLa cells. BRIT J PHARMACOL 1999 Vol 127, pages 1777-1786 Whole document	18-23
Y		1-17, 24-26

[X]	Further documents are listed in the continuation of Box C.	[X]	See patent family annex.
* "A"	Special categories of cited documents:  document defining the general state of the art which is not considered	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"E"	to be of particular relevance earlier application or patent but published on or after the international filing date	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"L"	document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination
"O"	document referring to an oral disclosure, use, exhibition or other means	""	being obvious to a person skilled in the art
"P"	document published prior to the international filing date but later than the priority date claimed	"&"	document member of the same patent family
Date	of the actual completion of the international search	Date	of mailing of the international search report
22 A	pril 2005 (22-04-2005)	18 M	ay 2005 (18-05-2005)
	te and mailing address of the ISA/CA	Auth	orized officer
Place 50 V	adian Intellectual Property Office e du Portage I, C114 - 1st Floor, Box PCT Victoria Street	Debo	ora Fujimoto (819) 997-1855
	neau, Quebec K1A 0C9 imile No.: 001(819)953-2476	]	
I aco	####♥ 190.0 UU 1(01 <i>7173)3=2</i> 4 / U		

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В	Box No. I Nuc	leotide and/or amino acid sequence(s) (Continuation of item 1.b of the first sheet)
1.	. With regard to any claimed invention,	nucleotide and/or amino acid sequence disclosed in the international application and necessary to the the international search was carried out on the basis of:
	a. type of material	
	[X] a sequence	e listing
	[ ] table(s) re	elated to the sequence listing
	b. format of materia	1
	[X] on paper	
	[X] in electron	nic form
	c. time of filing/furn	nishing
	[X] contained	in the international application as filed.
	[X] filed toget	ther with the international application in electronic form
	[ ] furnished	subsequently to this Authority for the purposes of search.
2.	[X] In addition, in t	he case that more than one version or copy of a sequence listing and/or table relating thereto has been
	filed or furnishe application as f	ed, the required statements that the information in the subsequent or additional copies is identical to that in the iled or does not go beyond the application as filed, as appropriate, were furnished.
3.	Additional comments	

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Box	N	o. I	Observations where certain claims were found unsearchable (Continuation of item 2 of the first sheet)
This reas			national search report has not been established in respect of certain claims under Article 17(2)(a) for the following
1.	٢X	1	Claim Nos.: 1 to 26
1.	L*-		because they relate to subject matter not required to be searched by this Authority, namely:
			Although claims 1 to 26 encompass a method of treatment of the human/animal body which this Authority is not obliged to search under Rule 39.1(iv) of the PCT, the search has been carried out based on the alleged effects of the compounds referred to therein.
2	г	,	Claim Nea
2.	L	-	Claim Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3.	[	-	Claim Nos.: because they are dependant claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box	No	). I	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This	In:	terr	national Searching Authority found multiple inventions in this international application, as follows:
1.	ĺ	]	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2.	[	1	As all searchable claims could be searched without effort justifying additional fees, this Authority did not invite payment of additional fees.
3.	[	J	As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claim Nos. :
4.	[	]	No required additional search fees were timely paid by the applicant. Consequently, this international search report is
			restricted to the invention first mentioned in the claims; it is covered by claim Nos. :
			Remark on Protest [ ] The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
			[ ] The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
			No protest accompanied the payment of additional search fees.

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C (Continua	ation). DOCUMENTS CONSIDERED TO BE RELEVANT	
Category*	Citation of document, with indication, where appropriate, of the relevant	Relevant to claim No.
х	BERG RW et al  The means to an end of tumor cell resistance to chemotherapeutic drugs targeting thymidylate synthase: shoot the messenger.  CURR DRUG TARGETS 2002 Vol 3, pages 297-309  Whole document	18-24
Y		1-17, 25, 26
Y	BERG RW et al Tumor growth inhibition in vivo and G <sub>2</sub> /M cell cycle arrest induced by antisense oligodeoxynucleotide targeting thymidylate synthase.  J PHARMACOL EXP THER 2001 Vol 298(2), pages 477-484  Whole document	1-26
Y	FERGUSON PJ et al Antisense-induced down-regulation of thymidylate synthase and enhanced cytotoxicity of 5-FUdR in 5-FUdR-resistant HeLa cells. BRIT J PHARMACOL 2001 Vol 134, pages 1437-1446 Whole document	1-26
Y	SCHMITZ JC ET AL Effect of 2'-O-methyl antisense ORNs on expression of thymidylate synthase in human colon cancer RKO cells. NUCLEIC ACIDS RES 2001 Vol 29(2), pages 415-422 Whole document	1-26

Information on patent family members

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